

SEQUENCE LISTING

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<120> Vaccine Delivery System and Method of Production

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<140> 09/308,435

<141> 1999-05-19

<150> PCT/SE99/00582

<151> 1999-04-09

<150> SE 9801288-3

<151> 1998-04-14

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<170> PatentIn Ver. 2.1

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<213> Helicobacter pylori

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975

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1 5 10 15

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<223> lipid chains a and b attached respectively at positions 3 and 2 of propyl group attached to sulfhydryl of cysteine residue at position 31

<400> 25

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Leu Gly Ala Ser Val Val Ala Leu Leu Val Gly Leu Ala Gly Cys
20 25 30



(1993) J. Bacteriol. 175, 674-683. Reference is also made to P W Toole et al, Bacteriology Vol. 177, No. 21, Nov. 1995; and Jones, A.C., Logan, R.P., Foynes, S., Cockayne, A., Wren, B.W. and Penn, C.W., J. Bacteriol. 179 (17), 5643-5647 (1997) which concern HpaA proteins.

The Hpa A protein is expressed by all *H. pylori* strains tested, and antibodies created towards this protein do not cross-react with common endogenous human bacteria of other species or with selected human tissues including the gastric mucosa. Thus being a well conserved putative adhesin with immunogenic properties, the HpaA protein is useful both for the detection of *H. pylori* infections as well as for the manufacture of vaccine compositions. Table 1 shows a comparison of HpaA amino acid sequences derived from 4 different strains of *H. Pylori*. It can be seen from the table that the sequence is highly conserved amongst different strains.

Table 1

	Evans (8826)	MKTNGHFKDFAWKKCLLGTSVVALLVGCSPHIIETNEVALKLNYHPASEKVQALDEKILL (SEQ ID NO:5)
	GTC (J99)	MKTNGHFKDFAWKKCFLGASVVALLVGCSPHIIETNEVALKLNYHPASEKVQALDEKILL (SEQ ID NO:6)
	Trust (17874)	MKTNGHFKDFAWKKCLLGASVGALLVGCSPHIIETNEVALKLNYHPASEKVQALDEKILL (SEQ ID NO:7)
20	Penn (11637)	MRANNHFKDFAWKKCLLGASVVALLVGCSPHIIETNEVALKLNYHPASEKVQALDEKILL (SEQ ID NO:8)
	TIGR (26695)	MKANNHEKDFAWKKCLLGASVVALLVGCSPHIIETNEVALKLNYHPASEKVQALDEKILL (SEQ ID NO:9)
		•::*:********
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25	GTC (J99)	LRPAFQYSDNIAKEYENKFKNQTTLKVEEILQNQGYKVINVDSSDKDDFSFAQKKEGYLA (SEQ ID NO:11)
	Trust (17874)	LRPAFQYSDNIAKEYENKFKNQTVLKVEQILQNQGYKVINVDSSDKDDFSFAQKKEGYLA (SEQ ID NO:12)
	Penn (11637)	LRPAFQYSDNIAKEYENKFKNQTALKVEQILQNQGYKVISVDSSDKDDFSFAQKKEGYLA (SEQ ID NO:13)
	TIGR (26695)	LRPAFQYSDNIAKEYENKFKNQTALKVEQILQNQGYKVISVDSSDKDDLSFSQKKEGYLA (SEQ ID NO:14)
	11011 (20070)	
		•;••••••••••

Evans (8826)	VAMIGEIVLRPD?KRTIQKKSEPGLLFSTGLDKMEGVLIPAGFVKVTILEPMSGESLDSF	(SEQ	TD	NO:13)
Evallo (erra)	VAMNGEIVLRPDPKRTIQKKSEPGLLFSTGLDKMEGVLIPAGFVKVTILEPMSGESLDSF	(SEQ	ID	NO:16)
	VAMNGEIVLRPDPKRTIQKKSEPGLLFSTGLDKMEGVLIPAGFVKVTILEPMSGESLDSF	(SEQ	ID	NO:16)
	VAMNGEIVLRPDPKRTIQKKSEPGLLFSTGLDKMEGVLIPAGFIKVTILEPMSGESLDSF			
Penn (11637)	VAMNGEIVLRPDPKKTIQKKSEFGLLFSTGLDKALEGVBTTAGTTKVTTTT	\\		

	TIGR (26695)	VAMNGEIVLRPDPKRTIQKKSEPGLLFSTGLDKMEGVLIPAGFVKVTILEPMSGESLDSF (SEQ ID NO:16)
		*** *********************************
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5	GTC (J99)	TMDLSELDIQEKFLKTTHSSHSGGLVSTMVKGTDNSNDAIKSALNKIFASIMQEMDKKLT (SEQ ID NO:18)
	Trust (17874)	TMDLSELDIQEKFLKTTHSSHSGGLVSTMVKGTDNSNDAIKSALNKIFGSIMQEIDKKLT (SEQ ID NO:19)
	Penn (11637)	TMDLSELDIQEKFLKTTHSSHSGGLVSTMVKGTDNSNDAIKSALNKIFANIMQEIDKKLT (SEQ ID NO:20)
	TIGR (26695)	TMDLSELDIQEKFLKTTHSSHSGGLVSTMVKGTDNSNDAIKSALNKIFANIMQEIDKKLT (SEQ ID NO:20)

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	GTC (J99)	QRNLESYQKDAKELKNKRNR (SEQ ID NO:21)
	Trust (17874)	QKNLESYQKDAKELKGKRNR (SEQ ID NO:22)
	Penn (11637)	QKNLESYQKDAKELKGKRNR (SEQ ID NO:22)
15	TIGR (26695)	QKNLESYQKDAKELKGKRNR (SEQ ID NO:22)
		*

"*" at a certain position denotes an identical amino acid in all sequences

"." at a certain position denotes conserved amino acids (eg, amino acids of the same charge type such as lysine or arginine at a certain position).

Penn (11637)	DNA sequence deposited in Genbank under Accession No. X92502
Trust (17874)	DNA sequence deposited in Genbank under Accession No.U35455
Evans (8826)	DNA sequence deposited in Genbank under Accession No.X61574
TIGR (26695)	DNA sequence deposited under Accession No. AE000591
GTC (J99)	DNA sequence obtained in-house.

The strain names are indicated in brackets, strain 8826 being obtained from SWISS-PROT accession Q48264.

25

III

5

H (Lipid Chain a)
H (Lipid Chain b)

٧

5

10

15



(1993) J. Bacteriol. 175, 674-683. Reference is also made to P W Toole et al, Bacteriology Vol. 177, No. 21, Nov. 1995; and Jones, A.C., Logan, R.P., Foynes, S., Cockayne, A., Wren, B.W. and Penn, C.W., J. Bacteriol. 179 (17), 5643-5647 (1997) which concern HpaA proteins.

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Table 1

		/mm-1-1-1-1-1-1-1-1-1-1-1-1-1-1-1-1-1-1-
	Evans (8826)	MKTNGHFKDFAWKKCLLGISVVALLVGCSPHIIETNEVALKLNYHPASEKVQALDEKILL (SEQ ID NO:5)
	GTC (J99)	MKTNGHFKDFAWKKCFLGASVVALLVGCSPHIIETNEVALKLNYHPASEKVQALDEKILL (SEQ ID NO:6)
	Trust (17874)	MKTNGHFKDFAWKKCLLGASVGALLVGCSPHIIETNEVALKLNYHPASEKVQALDEKILL (SEQ ID NO:7)
20	Penn (11637)	MRANNHFKOFAWKKCLLGASVVALLVGCSPHIIETNEVALKLNYHPASEKVQALDEKILL (SEQ_ID_NO:8)
	TIGR (26695)	MKANNHFKDFAWKKCLLGASVVALLVGCSPHIIETNEVALKLNYHPASEKVQALDEKILL (SEQ ID NO:9)
		:::********
	Evans (8826)	LKPAFQYSDNIAKEYENKFKNQTTLKVEEILQNQGYKVINVDSSDKDDFSFAQKKEGYLA (SEQ ID NO:10)
25	GTC (J99)	LRPAFQYSDNIAKEYENKFKNQTTLKVEEILQNQGYKVINVDSSDKDDFSFAQKKEGYLA (SEQ ID NO:11)
25	GTC (J99) Trust (17874)	
25		LRPAFQYSDNĮAKEYENKFKNQTTLKVEEILQNQGYKVINVDSSDKDDFSFAQKKEGYLA (SEQ ID NO:11)
25	Trust (17874)	LRPAFQYSDNIAKEYENKFKNQTTLKVEEILQNQGYKVINVDSSDKDDFSFAQKKEGYLA (SEQ ID NO:11) LRPAFQYSDNIAKEYENKFKNQTVLKVEQILQNQGYKVINVDSSDKDDFSFAQKKEGYLA (SEQ ID NO:12)

Evans (8826)	VAMIGEIVLRPD?KRTIQKKSEPGLLFSTGLDKMEGVLIPAGFVKVTILEPMSGESLDSF	(SEQ ID NO:13)
Evallo (ecc.)	VAMNGEIVLRPDPKRTIQKKSEPGLLFSTGLDKMEGVLIPAGFVKVTILEPMSGESLDSF	(SEQ ID NO:16)
	VAMNGEIVLRPDPKRTIQKKSEPGLLFSTGLDKMEGVLIPAGFVKVTILEPMSGESLDSF	(SEQ ID NO:16)
I Luge (2	VAMNGEIVLRPDPKRTIQKKSEPGLLFSTGLDKMEGVLIPAGFIKVTILEPMSGESLDSF	
Pann (11637)	VAMNGETVLRPDPRRITORNSDEGGDESTGESTGESTGESTGESTGESTGESTGESTGESTGESTG	

	TIGR (26695)	VAMNGEIVLRPDPKRTIQKKSEPGLLFSTGLDKMEGVLIPAGFVKVTILEPMSGESLDSF (SEQ ID NO:16)
		*** ***********************************
	Evans (8826)	TMDLSELDIQEKFLKTTHSSHSGGLVSTMVKGTDNSNDAIKSALNKIFASIMQEMDKKLT (SEQ ID NO:18)
5	GTC (J99)	TMDLSELDIQEKFLKTTHSSHSGGLVSTMVKGTDNSNDAIKSALNKIFASIMQEMDKKLT (SEQ_ID_NO:18)
	Trust (17874)	TMDLSELDIQEKFLKTTHSSHSGGLVSTMVKGTDNSNDAIKSALNKIFGSIMQEIDKKLT (SEQ_ID_NO:19)
	Penn (11637)	TMDLSELDIQEKFLKTTHSSHSGGLVSTMVKGTDNSNDAIKSALNKIFANIMQEIDKKLT (SEQ ID NO:20)
	TIGR (26695)	TMDLSELDIQEKFLKTTHSSHSGGLVSTMVKGTDNSNDAIKSALNKIFANIMQEIDKKLT (SEQ_ID_NO:20)

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	Evans (8826)	QRNLESYQKDAKELKNKRNR (SEQ ID NO:21)
	GTC (J99)	QRNLESYQKDAKELKNKRNR (<u>SEQ_ID_NO:21</u>)
	Trust (17874)	QKNLESYQKDAKELKGKRNR (SEQ ID NO:22)
	Penn (11637)	QKNLESYQKDAKELKGKRNR (SEQ ID NO:22)
15	TIGR (26695)	QKNLESYQKDAKELKGKRNR (SEQ ID NO:22)
		*;************

"*" at a certain position denotes an identical amino acid in all sequences

"." at a certain position denotes conserved amino acids (eg, amino acids of the same charge type such as lysine or arginine at a certain position).

Penn (11637)	DNA sequence deposited in Genbank under Accession No. X92502
Trust (17874)	DNA sequence deposited in Genbank under Accession No.U35455
Evans (8826)	DNA sequence deposited in Genbank under Accession No.X61574
TIGR (26695)	DNA sequence deposited under Accession No. AE000591
GTC (J99)	DNA sequence obtained in-house.

The strain names are indicated in brackets, strain 8826 being obtained from SWISS-PROT accession Q48264.

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III

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